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-	449	topotecan	USPAT;	2002/07/18 09:30
			US-PGPUB;	
			EPO; JPO;	
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1- 1	26	topotecan same liposom\$	USPAT;	2002/07/15 11:57
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			IBM_TDB	
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İ			EPO; JPO;	
			DERWENT;	
1			IBM_TDB	
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			DERWENT;	
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1			EPO; JPO;	
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•	5	(topotecan and camptothecin) same liposom\$ and	USPAT;	2002/07/17 09:57
		sphingomyelin and cholesterol	US-PGPUB;	
		,	EPO; JPO;	
		·	DERWENT;	
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			DERWENT;	
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     8 Mar 22
                 TRCTHERMO no longer available
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NEWS 10 Mar 28
                 LIPINSKI/CALC added for property searching in REGISTRY
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NEWS 17 Apr 22 BIOSIS Gene Names now available in TOXCENTER
NEWS 18 Apr 22 Federal Research in Progress (FEDRIP) now available
NEWS 19
         Jun 03 New e-mail delivery for search results now available
NEWS 20
        Jun 10
                MEDLINE Reload
NEWS 21 Jun 10
                 PCTFULL has been reloaded
NEWS 22 Jul 02 FOREGE no longer contains STANDARDS file segment
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NEWS EXPRESS
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              AND CURRENT DISCOVER FILE IS DATED 05 FEBRUARY 2002
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=> camptothecin

L1 6159 CAMPTOTHECIN

=> liposmom? and l1

0 LIPOSMOM? AND L1

=> liposmom?

L3 0 LIPOSMOM?

=> liposom?

L4 63645 LIPOSOM?

=> 14 and 11

L5 125 L4 AND L1

=> 15 and sphingomyelin

7 L5 AND SPHINGOMYELIN

=> 16 and topotecan

3 L6 AND TOPOTECAN

=> dis 17 ibib abs 1-3

ANSWER 1 OF 3 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:31220 CAPLUS

DOCUMENT NUMBER: 136:90962

TITLE: Improved liposomal camptothecins

and uses thereof

INVENTOR(S): Madden, Thomas D.; Semple, Sean C. PATENT ASSIGNEE(S): Inex Pharmaceuticals Corporation, Can.

SOURCE: PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ----------

WO 2002002078 A2 20020110 WO 2001-CA981 20010629 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,

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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
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      AU 2001070413
                         A5 20020114
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                                                                 20010629
PRIORITY APPLN. INFO .:
                                            US 2000-215556P P 20000630
                                            US 2001-264616P P 20010126
                                            WO 2001-CA981
                                                                 20010629
      This invention relates to improved liposomal
AΒ
      camptothecin compns. and methods of using such compns. for
      treating neoplasia and for inhibiting angiogenesis. The compns. and
      methods are useful for modulating the plasma circulation half-life of an
      active agent. Topotecan (I) was encapsulated in
      sphingomyelin: cholesterol (55:45, mol/mol) liposomes
      using Mg-A-2318 ionophore method. The initial drug-to-lipid ratio was 0.1
      (wt./wt.) and drug loading was typically 95-100%. The therapeutic index
     of the liposomes contg. I was 18 fold the free I in human breast
     cancer model MX-1.
L7
     ANSWER 2 OF 3 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER:
                           2002:31219 CAPLUS
DOCUMENT NUMBER:
                           136:90961
TITLE:
                           Liposomal antineoplastic drugs and uses
                           thereof
INVENTOR(S):
                           Madden, Thomas D.; Semple, Sean C.; Ahkong, Quet F.
PATENT ASSIGNEE(S):
                           Inex Pharmaceuticals Corporation, Can.
SOURCE:
                           PCT Int. Appl., 47 pp.
                           CODEN: PIXXD2
DOCUMENT TYPE:
                           Patent
LANGUAGE:
                           English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
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     WO 2002002077
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                                             WO 2001-CA925 20010629
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     AU 2001070385
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PRIORITY APPLN. INFO.:
                                           US 2000-215556P P
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                                           US 2001-264616P P 20010126
                                           WO 2001-CA925
                                                              W 20010629
     This invention relates to liposomal antineoplastic agents (e.g.,
AB
     camptothecin) compns. and methods of using such compns. for
     treating neoplasia and for inhibiting angiogenesis. The compns. and
     methods are useful for modulating the plasma circulation half-life of an
     active agent. Topotecan (I) was encapsulated in
     sphingomyelin:cholesterol (55:45, mol/mol) liposomes
     using Mg-A-2318 ionophore method. The initial drug-to-lipid ratio was 0.1
     (wt./wt.) and drug loading was typically 95-100%. The therapeutic index
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of the  ${f liposomes}$  contg. I was 18 fold the free I in human breast

cancer model MX-1.

ANSWER 3 OF 3 CAPLUS COPYRIGHT 2002 ACS L7 2000:277830 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

132:313695

TITLE:

Liposome-entrapped topoisomerase inhibitors

INVENTOR(S):

Slater, James Lloyd; Colbern, Gail T.; Working, Peter

PATENT ASSIGNEE(S):

Alza Corporation, USA PCT Int. Appl., 50 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                 KIND DATE
                                         APPLICATION NO. DATE
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                                       WO 1999-US24228 19991015
    WO 2000023052
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                     A1 20010808 EP 1999-954971
                                                          19991015
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    US 6355268
                           20020312
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                      В1
                                                          19991015
    NO 2001001844
                      А
                           20010410
                                         NO 2001-1844
                                                          20010410
PRIORITY APPLN. INFO.:
                                      US 1998-104671P P 19981016
                                      WO 1999-US24228 W 19991015
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A compn. for the treatment of tumor contg. a therapeutically ED of a AB topoisomerase I inhibitor or topoisomerase I/II inhibitor is described. The compn. includes liposomes having an outer surface and an inner surface defining aq. liposome compartment, and being composed of a vesicle-forming lipid and of a vesicle-forming lipid derivatized with a hydrophilic polymer to form a coating of hydrophilic polymer chains on both the inner and outer surfaces of the liposomes. Entrapped in the liposomes is the topoisomerase inhibitor at a concn. of at least about 0.10 .mu.M drug per .mu.N lipid. A topoisomerase I inhibitor is selected from the group consisting of camptothecin and derivs. thereof. 6

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT => liposom? and cholesterol

L8 10812 LIPOSOM? AND CHOLESTEROL

=> 18 and topotecan

L9 11 L8 AND TOPOTECAN

=> 19 and ionophore

L10 4 L9 AND IONOPHORE

=> dis 110 ibib abs

L10 ANSWER 1 OF 4 MEDLINE

ACCESSION NUMBER: 2000393094 MEDLINE

DOCUMENT NUMBER: 20365399 PubMed ID: 10910044

TITLE: Liposomal encapsulation of topotecan

enhances anticancer efficacy in murine and human xenograft

models.

AUTHOR: Tardi P; Choice E; Masin D; Redelmeier T; Bally M; Madden T

,

CORPORATE SOURCE: Inex Pharmaceuticals Corporation, Burnaby, British

Columbia, Canada.

SOURCE: CANCER RESEARCH, (2000 Jul 1) 60 (13) 3389-93.

Journal code: 2984705R. ISSN: 0008-5472.

PUB. COUNTRY: United States

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200008

ENTRY DATE: Entered STN: 20000824

Last Updated on STN: 20000824 Entered Medline: 20000817

AB Topotecan was encapsulated in sphingomyelin/cholesterol liposomes using an ionophore-generated proton gradient.

After i.v. injection, liposomal topotecan was eliminated from the plasma much more slowly than free drug, resulting in a 400-fold increase in plasma area under the curve. Further, high-performance liquid chromatography analysis of plasma samples demonstrated that topotecan was protected from hydrolysis within the liposomal carrier with >80% of the drug remaining as the active, lactone species up to 24 h. The improved pharmacokinetics observed with liposomal topotecan correlated with increased efficacy in both murine and human tumor models. In the L1210 ascitic tumor model, optimal doses of liposomal topotecan resulted in a 60-day survival rate of 60-80%, whereas in a L1210 liver metastasis model, 100% long-term survival (>60 days) was achieved. In contrast, long-term survivors were rarely seen after treatment with free topotecan. Further, in a human breast carcinoma model (MDA 435/LCC6), liposomal topotecan provided greatly improved increase in life span relative to the free drug. These results suggest that liposomal encapsulation can significantly enhance

=> dis 110 ibib abs 1-20

L10 ANSWER 1 OF 4 MEDLINE

ACCESSION NUMBER: 2000393094 MEDLINE

DOCUMENT NUMBER: 20365399 PubMed ID: 10910044

the therapeutic activity of topotecan.

TITLE: Liposomal encapsulation of topotecan

enhances anticancer efficacy in murine and human xenograft

models.

AUTHOR: Tardi P; Choice E; Masin D; Redelmeier T; Bally M; Madden T

D

CORPORATE SOURCE: Inex Pharmaceuticals Corporation, Burnaby, British

Columbia, Canada.

SOURCE: CANCER RESEARCH, (2000 Jul 1) 60 (13) 3389-93.

Journal code: 2984705R. ISSN: 0008-5472.

PUB. COUNTRY: United States

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200008

ENTRY DATE: Entered STN: 20000824

Last Updated on STN: 20000824 . Entered Medline: 20000817

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L10 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:31220 CAPLUS

DOCUMENT NUMBER: 136:90962

TITLE: Improved liposomal camptothecins and uses

thereof

INVENTOR(S): Madden, Thomas D.; Semple, Sean C.
PATENT ASSIGNEE(S): Inex Pharmaceuticals Corporation, Can.

SOURCE: PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent
LANGUAGE: English

LANGUAGE: English FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2002002078 A2 20020110 WO 2001-CA981 20010629

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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
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             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     AU 2001070413
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                                                           20010629
PRIORITY APPLN. INFO.:
                                        US 2000-215556P P 20000630
                                        US 2001-264616P P 20010126
                                        WO 2001-CA981
                                                         W 20010629
     This invention relates to improved liposomal camptothecin
ΑB
     compns. and methods of using such compns. for treating neoplasia and for
     inhibiting angiogenesis. The compns. and methods are useful for
     modulating the plasma circulation half-life of an active agent.
     Topotecan (I) was encapsulated in sphingomyelin:
     cholesterol (55:45, mol/mol) liposomes using Mq-A-2318
     ionophore method. The initial drug-to-lipid ratio was 0.1
     (wt./wt.) and drug loading was typically 95-100%. The therapeutic index
     of the liposomes contg. I was 18 fold the free I in human breast
     cancer model MX-1.
L10 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER:
                         2002:31219 CAPLUS
DOCUMENT NUMBER:
                         136:90961
TITLE:
                         Liposomal antineoplastic drugs and uses
                         thereof
INVENTOR(S):
                         Madden, Thomas D.; Semple, Sean C.; Ahkong, Quet F.
PATENT ASSIGNEE(S):
                         Inex Pharmaceuticals Corporation, Can.
SOURCE:
                         PCT Int. Appl., 47 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
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                     KIND DATE
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ΑB
     This invention relates to liposomal antineoplastic agents (e.g.,
     camptothecin) compns. and methods of using such compns. for treating
     neoplasia and for inhibiting angiogenesis. The compns. and methods are
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     Topotecan (I) was encapsulated in sphingomyelin:
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    ionophore method. The initial drug-to-lipid ratio was 0.1
     (wt./wt.) and drug loading was typically 95-100%. The therapeutic index
    of the liposomes contg. I was 18 fold the free I in human breast
    cancer model MX-1.
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## 09702165

ACCESSION NUMBER:

2000:494205 CAPLUS

DOCUMENT NUMBER:

133:212986

TITLE:

Liposomal encapsulation of topotecan

enhances anticancer efficacy in murine and human

xenograft models

AUTHOR(S):

Tardi, Paul; Choice, Edward; Masin, Dana; Redelmeier,

Thomas; Bally, Marcel; Madden, Thomas D.

CORPORATE SOURCE:

Department of Advanced Therapeutics, British Columbia

Cancer Agency, Vancouver, BC, V5Z 4E3, Can. Cancer Research (2000), 60(13), 3389-3393

CODEN: CNREA8; ISSN: 0008-5472

PUBLISHER:

SOURCE:

American Association for Cancer Research

DOCUMENT TYPE:

Journal English

LANGUAGE:

AB

Topotecan was encapsulated in sphingomyelin/cholesterol liposomes using an ionophore-generated proton gradient.

After i.v. injection, liposomal topotecan was

eliminated from the plasma much more slowly than free drug, resulting in a 400-fold increase in plasma area under the curve. Further,

high-performance liq. chromatog. anal. of plasma samples demonstrated that

topotecan was protected from hydrolysis within the

liposomal carrier with >80% of the drug remaining as the active,

lactone species up to 24 h. The improved pharmacokinetics obsd. with

liposomal topotecan correlated with increased efficacy

in both murine and human tumor models. In the L1210 ascitic tumor model, optimal doses of liposomal topotecan resulted in a

60-day survival rate of 60-80%, whereas in a L1210 liver metastasis model, 100% long-term survival (>60 days) was achieved. In contrast, long-term survivors were rarely seen after treatment with free topotecan.

Further, in a human breast carcinoma model (MDA 435/LCC6),

liposomal topotecan provided greatly improved increase

30

in life span relative to the free drug. These results suggest that liposomal encapsulation can significantly enhance the therapeutic

activity of topotecan.

REFERENCE COUNT:

THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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